

SEQUENCE LISTING

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Hasegawa, Mamoru

<120> Methods of Treating Inflammatory Diseases
Associated With Bone Destruction (as amended)

<130> 50026/055001

<150> PCT/JP2004/002887

<151> 2004-03-05

<150> JP 2003-075964

<151> 2003-03-19

<160> 6

<170> PatentIn version 3.1

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<211> 900

<212> DNA

<213> Homo sapiens

<220>

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Thr Leu Cys Thr Ala Arg Pro Ser Pro Thr Leu Pro Glu Gln Asp Ala	
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ctc ccc tcc tcg gag gat gat gat gat gat gac tcc tct tca gag	144
Leu Pro Ser Ser Glu Asp Asp Asp Asp Asp Asp Ser Ser Ser Glu	
35 40 45	

gag aaa gaa aca gat aac acc aaa cca aac ccc gta gct cca tat tgg	192
Glu Lys Glu Thr Asp Asn Thr Lys Pro Asn Pro Val Ala Pro Tyr Trp	
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aca tcc cca gaa aag atg gaa aag aaa ttg cat gca gtg ccg gct gcc	240
Thr Ser Pro Glu Lys Met Glu Lys Lys Leu His Ala Val Pro Ala Ala	
65 70 75 80	

aag aca gtg aag ttc aaa tgc cct tcc agt ggg acc cca aac ccc aca	288
Lys Thr Val Lys Phe Lys Cys Pro Ser Ser Gly Thr Pro Asn Pro Thr	
85 90 95	

ctg cgc tgg ttg aaa aat ggc aaa gaa ttc aaa cct gac cac aga att	336
Leu Arg Trp Leu Lys Asn Gly Lys Glu Phe Lys Pro Asp His Arg Ile	
100 105 110	
gga ggc tac aag gtc cgt tat gcc acc tgg agc atc ata atg gac tct	384
Gly Gly Tyr Lys Val Arg Tyr Ala Thr Trp Ser Ile Ile Met Asp Ser	
115 120 125	
gtg gtg ccc tct gac aag ggc aac tac acc tgc att gtg gag aat gag	432
Val Val Pro Ser Asp Lys Gly Asn Tyr Thr Cys Ile Val Glu Asn Glu	
130 135 140	
tac ggc agc atc aac cac aca tac cag ctg gat gtc gtg gag cgg tcc	480
Tyr Gly Ser Ile Asn His Thr Tyr Gln Leu Asp Val Val Glu Arg Ser	
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cct cac cgg ccc atc ctg caa gca ggg ttg ccc gcc aac aaa aca gtg	528
Pro His Arg Pro Ile Leu Gln Ala Gly Leu Pro Ala Asn Lys Thr Val	
165 170 175	
gcc ctg ggt agc aac gtg gag ttc atg tgt aag gtg tac agt gac ccg	576
Ala Leu Gly Ser Asn Val Glu Phe Met Cys Lys Val Tyr Ser Asp Pro	
180 185 190	
cag ccg cac atc cag tgg cta aag cac atc gag gtg aat ggg agc aag	624
Gln Pro His Ile Gln Trp Leu Lys His Ile Glu Val Asn Gly Ser Lys	
195 200 205	
att ggc cca gac aac ctg cct tat gtc cag atc ttg aag gta atc atg	672
Ile Gly Pro Asp Asn Leu Pro Tyr Val Gln Ile Leu Lys Val Ile Met	
210 215 220	
gca cca gtc ttc gtg ggc cag tct act ggg aag gag acc act gtc tcg	720
Ala Pro Val Phe Val Gly Gln Ser Thr Gly Lys Glu Thr Thr Val Ser	
225 230 235 240	
ggg gct caa gtt cct gtg ggc agg ctc agt tgc ccc cga atg gga tca	768
Gly Ala Gln Val Pro Val Gly Arg Leu Ser Cys Pro Arg Met Gly Ser	
245 250 255	
ttc ctc acg ctt cag gca cac aca ctc cat ctc agt agg gat cta gcc	816
Phe Leu Thr Leu Gln Ala His Thr Leu His Leu Ser Arg Asp Leu Ala	
260 265 270	
aca tcc ccc agg act agt aac aga ggt cac aaa gtg gag gtg agc tgg	864
Thr Ser Pro Arg Thr Ser Asn Arg Gly His Lys Val Glu Val Ser Trp	
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35 40 45

Glu Lys Glu Thr Asp Asn Thr Lys Pro Asn Pro Val Ala Pro Tyr Trp
50 55 60

Thr Ser Pro Glu Lys Met Glu Lys Lys Leu His Ala Val Pro Ala Ala
65 70 75 80

Lys Thr Val Lys Phe Lys Cys Pro Ser Ser Gly Thr Pro Asn Pro Thr
85 90 95

Leu Arg Trp Leu Lys Asn Gly Lys Glu Phe Lys Pro Asp His Arg Ile
100 105 110

Gly Gly Tyr Lys Val Arg Tyr Ala Thr Trp Ser Ile Ile Met Asp Ser
115 120 125

Val Val Pro Ser Asp Lys Gly Asn Tyr Thr Cys Ile Val Glu Asn Glu
130 135 140

Tyr Gly Ser Ile Asn His Thr Tyr Gln Leu Asp Val Val Glu Arg Ser
145 150 155 160

Pro His Arg Pro Ile Leu Gln Ala Gly Leu Pro Ala Asn Lys Thr Val
165 170 175

Ala Leu Gly Ser Asn Val Glu Phe Met Cys Lys Val Tyr Ser Asp Pro
180 185 190

Gln Pro His Ile Gln Trp Leu Lys His Ile Glu Val Asn Gly Ser Lys
195 200 205

Ile Gly Pro Asp Asn Leu Pro Tyr Val Gln Ile Leu Lys Val Ile Met
210 215 220

Ala Pro Val Phe Val Gly Gln Ser Thr Gly Lys Glu Thr Thr Val Ser
225 230 235 240

Gly Ala Gln Val Pro Val Gly Arg Leu Ser Cys Pro Arg Met Gly Ser
245 250 255

Phe Leu Thr Leu Gln Ala His Thr Leu His Leu Ser Arg Asp Leu Ala
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 Thr Pro Arg Asp Gly Gly Arg Gln Arg Gly Glu Pro Asp Pro Arg Asp
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 atc cga aac acc aat gag tac aca gag ggg cct act gtc gtc cca aga 192
 Ile Arg Asn Thr Asn Glu Tyr Thr Glu Gly Pro Thr Val Val Pro Arg
 50 55 60
 cct ggg ctc aag cct gct cct cgc ccc tcc act cag cac aaa cac gag 240
 Pro Gly Leu Lys Pro Ala Pro Arg Pro Ser Thr Gln His Lys His Glu
 65 70 75 80
 aga ctc cac ggt ctg cct gag cac cgc cag cct cct agg ctc cag cac 288
 Arg Leu His Gly Leu Pro Glu His Arg Gln Pro Pro Arg Leu Gln His
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 tcg cag gtc cat tct tct gca cga gcc cct ctg tcc aga tcc ata agc 336
 Ser Gln Val His Ser Ser Ala Arg Ala Pro Leu Ser Arg Ser Ile Ser
 100 105 110
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 Thr Val Ser Ser Gly Ser Arg Ser Ser Thr Arg Thr Ser Thr Ser Ser
 115 120 125
 agc tcc tct gaa cag aga ctg cta gga tca tcc ttc tcc tcc ggg cct 432
 Ser Ser Ser Glu Gln Arg Leu Leu Gly Ser Ser Phe Ser Ser Gly Pro
 130 135 140
 gtt gct gat ggc ata atc cgg gtg caa ccc aaa tct gag ctc aag cca 480
 Val Ala Asp Gly Ile Ile Arg Val Gln Pro Lys Ser Glu Leu Lys Pro
 145 150 155 160
 ggt gag ctt aag cca ctg agc aag gaa gat ttg ggc ctg cac gcc tac 528
 Gly Glu Leu Lys Pro Leu Ser Lys Glu Asp Leu Gly Leu His Ala Tyr

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Arg Cys Glu Asp Cys Gly Lys Cys Lys Cys Lys Glu Cys Thr Tyr Pro				
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Arg Pro Leu Pro Ser Asp Trp Ile Cys Asp Lys Gln Cys Leu Cys Ser				
	195	200	205	
gcc cag aac gtg att gac tat ggg act tgt gta tgc tgt gtg aaa ggt				672
Ala Gln Asn Val Ile Asp Tyr Gly Thr Cys Val Cys Cys Val Lys Gly				
	210	215	220	
ctc ttc tat cac tgt tct aat gat gat gag gac aac tgt gct gac aac				720
Leu Phe Tyr His Cys Ser Asn Asp Asp Glu Asp Asn Cys Ala Asp Asn				
	225	230	235	240
cca tgt tct tgc agc cag tct cac tgt tgt aca cga tgg tca gcc atg				768
Pro Cys Ser Cys Ser Gln Ser His Cys Cys Thr Arg Trp Ser Ala Met				
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Lys Gly Cys Leu Lys Leu Cys Gln Gly Cys Tyr Asp Arg Val Asn Arg				
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Ile Arg Asn Thr Asn Glu Tyr Thr Glu Gly Pro Thr Val Val Pro Arg			
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 Arg Leu His Gly Leu Pro Glu His Arg Gln Pro Pro Arg Leu Gln His
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 Thr Val Ser Ser Gly Ser Arg Ser Ser Thr Arg Thr Ser Thr Ser Ser
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 Ser Ser Ser Glu Gln Arg Leu Leu Gly Ser Ser Phe Ser Ser Gly Pro
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 145 150 155 160
 Gly Glu Leu Lys Pro Leu Ser Lys Glu Asp Leu Gly Leu His Ala Tyr
 165 170 175
 Arg Cys Glu Asp Cys Gly Lys Cys Lys Cys Lys Glu Cys Thr Tyr Pro
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 Arg Pro Leu Pro Ser Asp Trp Ile Cys Asp Lys Gln Cys Leu Cys Ser
 195 200 205
 Ala Gln Asn Val Ile Asp Tyr Gly Thr Cys Val Cys Cys Val Lys Gly
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 Leu Phe Tyr His Cys Ser Asn Asp Asp Glu Asp Asn Cys Ala Asp Asn
 225 230 235 240
 Pro Cys Ser Cys Ser Gln Ser His Cys Cys Thr Arg Trp Ser Ala Met
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 Gly Val Met Ser Leu Phe Leu Pro Cys Leu Trp Cys Tyr Leu Pro Ala
 260 265 270
 Lys Gly Cys Leu Lys Leu Cys Gln Gly Cys Tyr Asp Arg Val Asn Arg
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Lys Ala Val Val Met Thr Arg Asp Asp Ser Ser Gly Gly Trp Phe Pro	
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cag gaa gga ggc ggg atc agt cgc gtc ggc gtg tgt aag gtc atg cac	144
Gln Glu Gly Gly Gly Ile Ser Arg Val Gly Val Cys Lys Val Met His	
35 40 45	
cct gaa ggc aac gga cga agc ggc ttt ctc atc cat ggc gag cga cag	192
Pro Glu Gly Asn Gly Arg Ser Gly Phe Leu Ile His Gly Glu Arg Gln	
50 55 60	
aaa gac aaa ctg gtg gta ttg gaa tgc tat gtc aga aag gac ttg gtc	240
Lys Asp Lys Leu Val Val Leu Glu Cys Tyr Val Arg Lys Asp Leu Val	
65 70 75 80	
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Tyr Thr Lys Ala Asn Pro Thr Phe His His Trp Lys Val Asp Asn Arg	
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Lys Phe Gly Leu Thr Phe Gln Ser Pro Ala Asp Ala Arg Ala Phe Asp	
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Arg Gly Val Arg Lys Ala Ile Glu Asp Leu Ile Glu Gly Ser Thr Thr	
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Ser Ser Ser Thr Leu His Asn Glu Ala Glu Leu Gly Asp Asp Asp Val	
130 135 140	
ttc acg aca gct acg gac agt tct tct aat tcc tcg cag aag agg gag	480
Phe Thr Thr Ala Thr Asp Ser Ser Ser Asn Ser Ser Gln Lys Arg Glu	
145 150 155 160	
ccg act acg agg aca atc tcc tcc ccc acg tcc tgt gag cac cgg aag	528
Pro Thr Thr Arg Thr Ile Ser Ser Pro Thr Ser Cys Glu His Arg Lys	
165 170 175	
att tat acc ctt gac cca tac ccc atg gac cat tac cac cct gac cag	576
Ile Tyr Thr Leu Asp Pro Tyr Pro Met Asp His Tyr His Pro Asp Gln	
180 185 190	
cgg ttg ccg cgg tcc tac ccc cag gtc acc ttc cca gaa gat gat gaa	624
Arg Leu Pro Arg Ser Tyr Pro Gln Val Thr Phe Pro Glu Asp Asp Glu	
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gaa att gta cgc atc aac ccc cga gag aag atc tgg atg acc ggt tat	672

Glu Ile Val Arg Ile Asn Pro Arg Glu Lys Ile Trp Met Thr Gly Tyr	
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gaa gac tac cgg cac gcg ccg gtt cgc ggc aaa tac tta gac acc aca	720
Glu Asp Tyr Arg His Ala Pro Val Arg Gly Lys Tyr Leu Asp Thr Thr	
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gaa gac gcg gac tcc tac gtg cgc ttc gcc aag ggc gaa gtc ccc aaa	768
Glu Asp Ala Asp Ser Tyr Val Arg Phe Ala Lys Gly Glu Val Pro Lys	
245 250 255	
cac gaa tat acc tat ccc tat gtt gat tct tcg gac ttc ggc ttc ggg	816
His Glu Tyr Thr Tyr Pro Tyr Val Asp Ser Ser Asp Phe Gly Phe Gly	
260 265 270	
gag gat ccc aaa ggt agt gtg atc aag aca cag ccg ccc agg gcc aag	864
Glu Asp Pro Lys Gly Ser Val Ile Lys Thr Gln Pro Pro Arg Ala Lys	
275 280 285	
tcc cgt cgg cgg aag gag aac ggc gaa cgg tcg cgg tgt gtg tac tgc	912
Ser Arg Arg Arg Lys Glu Asn Gly Glu Arg Ser Arg Cys Val Tyr Cys	
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305 310 315 320	
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Ala Pro Asp Ala Val Arg Thr Cys Ile Arg Arg Val Ser Cys Met Trp	
325 330 335	
tgc gcg gac agc atg ctg tac cac tgt atg tcc gac ccc gag gga gac	1056
Cys Ala Asp Ser Met Leu Tyr His Cys Met Ser Asp Pro Glu Gly Asp	
340 345 350	
tac act gac cct tgt tcg tgt gac aca agc gat gag aag ttt tgc ctc	1104
Tyr Thr Asp Pro Cys Ser Cys Asp Thr Ser Asp Glu Lys Phe Cys Leu	
355 360 365	
cgg tgg atg gct cta att gcc ttg tct ttc ctg gcc cct tgt atg tgc	1152
Arg Trp Met Ala Leu Ile Ala Leu Ser Phe Leu Ala Pro Cys Met Cys	
370 375 380	
tgt tac ctg ccc ctc cgg gcc tgc cac cgc tgt gga gtg atg tgc agg	1200
Cys Tyr Leu Pro Leu Arg Ala Cys His Arg Cys Gly Val Met Cys Arg	
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Cys Cys Gly Gly Lys His Lys Ala Ala	
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35 40 45

Pro Glu Gly Asn Gly Arg Ser Gly Phe Leu Ile His Gly Glu Arg Gln
50 55 60

Lys Asp Lys Leu Val Val Leu Glu Cys Tyr Val Arg Lys Asp Leu Val
65 70 75 80

Tyr Thr Lys Ala Asn Pro Thr Phe His His Trp Lys Val Asp Asn Arg
85 90 95

Lys Phe Gly Leu Thr Phe Gln Ser Pro Ala Asp Ala Arg Ala Phe Asp
100 105 110

Arg Gly Val Arg Lys Ala Ile Glu Asp Leu Ile Glu Gly Ser Thr Thr
115 120 125

Ser Ser Ser Thr Leu His Asn Glu Ala Glu Leu Gly Asp Asp Asp Val
130 135 140

Phe Thr Thr Ala Thr Asp Ser Ser Ser Asn Ser Ser Gln Lys Arg Glu
145 150 155 160

Pro Thr Thr Arg Thr Ile Ser Ser Pro Thr Ser Cys Glu His Arg Lys
165 170 175

Ile Tyr Thr Leu Asp Pro Tyr Pro Met Asp His Tyr His Pro Asp Gln
180 185 190

Arg Leu Pro Arg Ser Tyr Pro Gln Val Thr Phe Pro Glu Asp Asp Glu
195 200 205

Glu Ile Val Arg Ile Asn Pro Arg Glu Lys Ile Trp Met Thr Gly Tyr
210 215 220

Glu Asp Tyr Arg His Ala Pro Val Arg Gly Lys Tyr Leu Asp Thr Thr
225 230 235 240

Glu Asp Ala Asp Ser Tyr Val Arg Phe Ala Lys Gly Glu Val Pro Lys
245 250 255

His Glu Tyr Thr Tyr Pro Tyr Val Asp Ser Ser Asp Phe Gly Phe Gly
260 265 270

Glu Asp Pro Lys Gly Ser Val Ile Lys Thr Gln Pro Pro Arg Ala Lys
275 280 285

Ser Arg Arg Arg Lys Glu Asn Gly Glu Arg Ser Arg Cys Val Tyr Cys

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Arg Asp Met Phe Asn His Glu Glu Asn Arg Arg Gly His Cys Gln Asp				
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Ala Pro Asp Ala Val Arg Thr Cys Ile Arg Arg Val Ser Cys Met Trp				
	325		330	335
Cys Ala Asp Ser Met Leu Tyr His Cys Met Ser Asp Pro Glu Gly Asp				
	340		345	350
Tyr Thr Asp Pro Cys Ser Cys Asp Thr Ser Asp Glu Lys Phe Cys Leu				
	355		360	365
Arg Trp Met Ala Leu Ile Ala Leu Ser Phe Leu Ala Pro Cys Met Cys				
	370		375	380
Cys Tyr Leu Pro Leu Arg Ala Cys His Arg Cys Gly Val Met Cys Arg				
385		390		395 400
Cys Cys Gly Gly Lys His Lys Ala Ala Ala				
	405		410	